U.S. PATENT AND TRADEMARK OFFICE; U.S. DEPARTMENT OF COMMERC FORM PTO-1390 (Modified) REV. 2-2005) 004974.01070 TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) U.S. APPLICATION NO. (If known, see 37 CFR 1.5) 0/550327 CONCERNING A SUBMISSION UNDER 35 U.S.C. 371 INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED 17 March 2004 28 March 2003 PCT/EP2004/002731 TITLE OF INVENTION DIAGNOSTICS AND THERAPEUTICS FOR DISEASES ASSOCIATED WITH ENDOTHELIAL DIFFERENTIATION, SPHINGOLIPID G-PROTEIN-COUPLED RECEPTOR 6(EDG6) APPLICANT(S) FOR DO/EO/US GOLZ, Stefan et al Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: × This is a FIRST submission of items concerning a submission under 35 U.S.C. 371. This is a SECOND or SUBSEQUENT submission of items concerning a submission under 35 U.S.C. 371. 2. Ø This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), 3. (9) and (24) indicated below. 4. \bowtie The US has been elected (Article 31). A copy of the International Application as filed (35 U.S.C. 371 (c)(2)) is attached hereto (required only if not communicated by the International Bureau). b. 🖾 has been communicated by the International Bureau. is not required, as the application was filed in the United States Receiving Office (RO/US). c. 🗆 An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)). 6. a. 🗆 is attached hereto. has been previously submitted under 35 U.S.C. 154(d)(4). b. \Box Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371 (c)(3)) 7. are attached hereto (required only if not communicated by the International Bureau). b. 🗆 have been communicated by the International Bureau. c. 🗆 have not been made; however, the time limit for making such amendments has NOT expired. d. 🗆 have not been made and will not be made. An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)). 8. An oath or declaration of the inventor(s) (35 U.S.C. 371 (c)(4)). 9. An English language translation of the annexes to the International Preliminary Examination Report under PCT 10. Article 36 (35 U.S.C. 371 (c)(5)). A copy of the International Preliminary Examination Report (PCT/IPEA/409). 11. A copy of the International Search Report (PCT/ISA/210). 12. \boxtimes Items 13 to 23 below concern document(s) or information included: 13. An Information Disclosure Statement under 37 CFR 1.97 and 1.98. An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. 14 15. A FIRST preliminary amendment. A SECOND or SUBSEQUENT preliminary amendment. 16 17. A substitute specification. 18 A power of attorney and/or change of address letter.

A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 - 1.825.

A second copy of the English language translation of the International Application under 35 U.S.C. 154(d)(4).

A second copy of the published International Application under 35 U.S.C. 154(d)(4).

Transmittal of Computer-Readable Form of Sequence Listing and ADS

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Express Mail Label No.

Other items or information:

JC14 Rec'd PCT/PTO 22 SEP 2005

Approved for use through 3/31/2007. OMB 0651-0201

U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number. INTERNATIONAL APPLICATION NO. U.S. APPLICATION NO (if known, see 37 CFR 1.5) ATTORNEY'S DOCKET NUMBER 004974.01070 PCT/EP2004/002731 **CALCULATIONS PTO USE** The following fees are submitted: 24. \$300 \$300.00 25. Examination fee If International preliminary examination report prepared by USPTO and all claims \$100 \$200.00 satisfy provisions of PCT Article 33(1)-(4)..... \$200 Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to \$100 \$400 \$400.00 \$500 \$900.00 TOTAL OF 24, 25 and 26 = Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing or computer program listing filed in an electronic medium). The fee is \$250 for each additional 50 sheets of paper or fraction thereof. Number of each additional 50 or RATE **Total Sheets** Extra Sheets fraction thereof (round up to a whole /50 = \$250.00 \$ \$250.00 - 100 = 136 36 Surcharge of \$130.00 for furnishing the oath or declaration later than months from the \$ earliest claimed priority date (37 CFR 1.492(e)). NUMBER EXTRA RATE **CLAIMS** NUMBER FILED - 20 = \$50.00 \$50.00 Total claims 21 х 1 Independent claims 8 5 \$200.00 \$ \$1,000.00 х \Box MULTIPLE DEPENDENT CLAIMS (if applicable) \$360.00 \$ \$0.00 TOTAL OF ABOVE CALCULATIONS = \$ \$2,200.00 Applicant claims small entity status. See 37 CFR 1.27. The fees indicated above are \$ \$0.00 reduced by 1/2. SUBTOTAL = \$ \$2,200.00 Processing fee of \$130.00 for furnishing the English translation later than 30 months from \$ \$0.00 the earliest claimed priority date (37 CFR 1.492(f)). \$ \$2,200.00 **TOTAL NATIONAL FEE =** Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be \$ \$0.00 accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40 per property \$2,200.00 **TOTAL FEES ENCLOSED =** Amount to be \$ Amount to be \$ a. A check in the amount of \$ to cover the above fees is enclosed. \$2,200.00 to cover the above fees. 19-0733 in the amount of Please charge my Deposit Account No. A duplicate copy of this sheet is enclosed. The Director is hereby authorized to charge any additional fees which may be required, or credit any overpayment . A duplicate copy of this sheet is enclosed. to Deposit Account No. 19-0733 Fees are to be charged to a credit card. WARNING: Information on this form may become public. Credit card d. 🔲 information should not be included on this form. Provide credit card information and authorization on PTO-2038. NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the International Application to pending status. Loss U. Hennerdinger SEND ALL CORRESPONDENCE TO: Banner & Witcoff, Ltd. Lisa M. Hemmendinger **Customer Number: 22907** NAME 42,653 Date Sepkulses 22,2005 REGISTRATION NUMBER